DOCKET NO.: ISIS-5027

PATENT

deoxy-3'-amino phosphoramidate, 5'-deoxy-5'-amino phosphoramidate, 3'-deoxy-3'-thio phosphorothioate, 5'-deoxy-5'-thio phosphorothioate or hydrogen phosphonate linkage.

REMARKS

Applicant submitted a Preliminary Amendment on February 20, 2002, canceling claim 1 and adding new claims 94-154. Applicant presents herein a Second Preliminary Amendment in which new claims 155-164 have been added. New claims 155-164 find support throughout the specification and claims as filed, including, for example, in the specification at pages 4, 6-14, 17, 18, 20-27, 30, 32, 91, 92, and 94-96, Figure 4, and claim 1 as originally filed.

Upon entry of this amendment, claims 94-164 will be pending.

Attached hereto is a marked-up version of the changes made to the specification and claims by the current amendment. The attached page is captioned "Version with markings to show changes made."

Applicant respectfully requests that this amendment be entered without prejudice.

Applicant believes all of the claims presently before the Examiner patentably define the invention over the prior art and are otherwise in condition for ready allowance. Applicant respectfully requests early notification of the same.

Respectfully submitted,

Date: April 1,2002

Gwilym J.O. Attwell Registration No. 45,449

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VERSION WITH MARKINGS TO SHOW CHANGES MADE

In the Claims:

Please add new claims 155-164.